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# Prevalence, trend, and predictor analyses of vitamin D deficiency in the US population, 2001–2018

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**Background:** The National Health and Nutrition Examination Surveys (NHANES) collect and release data to the public every 2 years. The latest NHANES study on the vitamin D status of Americans was based on data from 2001 to 2014, and the latest data (2015–2016 and 2017–2018) have not been studied yet. Thus, we extracted all the available data from NHANES (2001–2018), aiming to analyze the prevalence and trends of vitamin D deficiency (VDD) in the US population to bridge the research gap.

**Methods:** According to previous studies and nutritional guidelines for vitamin D, severe VDD was defined as serum 25(OH)D levels of <25 nmol/L, moderate deficiency as 25–50 nmol/L, insufficiency as 50–75 nmol/L, and sufficiency as >75 nmol/L. We comprehensively estimated the prevalence of serum 25(OH)D levels of <25, 25–50, 50–75, and >75 nmol/L in Americans and described trends in vitamin D status from 2001 to 2018. Weighted multivariate linear regression models were used to explore the predictors of VDD. All analyses and the data were adjusted for the complex sampling design of NHANES using Mobile Examination Center (MEC) weights.

**Results:** Based on the most recent data of 71,685 participants, our study showed that the weighted prevalence of severe and moderate VDD was 2.6% and 22.0%, and the prevalence of vitamin D insufficiency (VDI) and sufficiency was 40.9% and 34.5%. The prevalence of severe and moderate VDD was higher in women, non-Hispanic black Americans, people aged 20–29 years, and during the season of winter. From 2001 to 2018, we found a slight linear decrease in the prevalence of moderate VDD (coefficient =  $-0.847$ ;  $P = 0.009$ ) and VDI (coefficient =  $-0.810$ ;  $P = 0.014$ ). We also found a slight linear increase in vitamin D sufficient (coefficient =  $1.693$ ;  $P = 0.004$ ). However, no trend change was observed in severe VDD (coefficient =  $-0.037$ ;  $P = 0.698$ ). Age, sex, ethnicity, season, sun-protective behaviors, lower BMI, lower socioeconomic status (SES), drinking, and lower milk consumption were predictors of severe VDD.

**Conclusion:** Vitamin D deficiency is still prevalent in the United States, especially in non-Hispanic black Americans, women, individuals aged

20–29, and during winter. Therefore, individuals, healthcare providers, and policymakers should take public health measures to develop and implement prevention strategies to deal with VDD.

#### KEYWORDS

prevalence, trends, vitamin D, predictors, NHANES

## Introduction

Vitamin D plays a critical role in maintaining bone mineralization by influencing calcium and phosphorus homeostasis (1). Vitamin D deficiency (VDD) is associated with an increased risk of rickets and osteomalacia (2, 3). Evidence has shown that VDD is associated with extraskeletal conditions, such as infection, cancer, diabetes mellitus, cardiovascular disease, and autoimmune disease (4–7). Recently, Pugach et al. (8) reported a strong association between vitamin D status and the mortality rate from COVID-19 in Europe. Vitamin D deficiency has become a global health issue (9), which is usually linked to insufficient exposure to sunlight and insufficient consumption of vitamin D-rich foods (10). Ultraviolet-B (UVB) radiation (wavelength 290–315 nm) in the sun rays could penetrate the skin and convert 7-dehydrocholesterol to pre-vitamin D<sub>3</sub> and then vitamin D<sub>3</sub> (11). *In vivo*, vitamin D absorbed from food is hydroxylated in the liver to 25-hydroxyvitamin D, then in the kidneys to 1,25-dihydroxyvitamin D to play a fundamental role in bone metabolism or other physiological processes. The Institute of Medicine (IOM) recommended 600 IU/d vitamin D intake for ages 1–70 years and 800 IU/d for ages 71 years and older (12). A recent study showed that the prevalence of VDD was still high in sunny Africa, and 32.2% of Africans were estimated to have VDD when a cut-off of serum 25(OH)D <50 nmol/L was used (13). In the United States, VDD is also a growing public health concern (14). However, due to the controversial definition of VDD, the results of different studies varied widely (12, 15).

Vitamin D deficiency has been defined differently depending on the effects on parathyroid hormone suppression, maximum calcium absorption, or bone mineral density (BMD) (16). Institute of Medicine defined people at risk of VDD as serum 25(OH)D < 30 nmol/L (12 ng/L) and at risk of insufficiency as 30 <25(OH)D <50 nmol/L (12–20 ng/L) (12). However, the Endocrine Society (ES) suggested VDD as a serum 25(OH)D <50 nmol/L and vitamin D insufficiency (VDI) as 50 <25(OH)D <75 nmol/L (17). Several National Health and Nutrition Examination Survey (NHANES) studies have estimated the vitamin D status of Americans by different cut-offs. National Health and Nutrition Examination Survey (2000–2004) reported that 5% of Americans had 25(OH)D levels <27.5 nmol/L (15). Liu et al. (18) estimated that 28.9% and 41.4% of

American adults had VDD [25(OH)D <50 nmol/L] and VDI [50 ≤25(OH)D <75 nmol/L] using NHANES 2001–2011. In this study, the prevalence was much higher among the elderly, non-Hispanic black Americans, people with obesity, and people who reported little milk consumption. Another study by Herrick et al. (19) investigated the vitamin D status in America using NHANES 2011–2014. They found the prevalence of 25(OH)D levels <30 and 30–50 nmol/L was 5.0% and 18.3%, respectively.

In August 2021 and April 2022, the NHANES website released the data on vitamin D from 2015 to 2016 and 2017 to 2018. However, there has not been a study analyzing these data. Therefore, this study aims to examine the American population's vitamin D status using currently available data from NHANES (2001–2018). This report also provides the trends for VDD and VDI in the United States from 2001 to 2018.

## Methods

### Study design and population

National Health and Nutrition Examination Survey is a nationally representative nutrition survey of general populations in the United States using a stratified, multi-stage random sampling design. As the latest data for vitamin D status are available as of NHANES 2017–2018, nine consecutive cycles of NHANES (2001–2002, 2003–2004, 2005–2006, 2007–2008, 2009–2010, 2011–2012, 2013–2014, 2015–2016, and 2017–2018) were selected for our analysis. Serum 25(OH)D data were available for subjects older than 6 years in 2001–2002 and older than 1 year in 2003–2018. We initially included 91,351 subjects from NHANES 2001–2018. After excluding 19,666 subjects without serum 25(OH)D data, 71,685 eligible subjects were included in this analysis. 25(OH)D concentrations were measured in the south during the winter months (November–March) and in the north during the summer months (April–October). The total serum 25(OH)D concentration was defined as the sum of 25(OH)D<sub>2</sub> and 25(OH)D<sub>3</sub>. Serum 25(OH)D concentrations were measured at the National Center for Environmental Health using the DiaSorin RIA kit (Stillwater MN) in 2001–2006 and the standardized liquid chromatography-tandem mass spectrometry (LC-MS/MS) method in 2007–2018. Considering the quality control issues of the RIA in NHANES 2001–2006, the Centers for

Disease Control and Prevention (CDC) decided to adjust and convert the measured values of 25(OH)D concentrations to equivalent measurements by LC-MS/MS methods. It allows us to combine and compare these data. The detailed method for adjustment can be found on the NHANES website and in [Appendix 1 in Supplementary material](#).

Since the definition of VDD was still controversial, we defined severe VDD as serum 25(OH)D levels of <25 nmol/L, moderate deficiency as 25–50 nmol/L, insufficiency as 50–75 nmol/L, and sufficient as >75 nmol/L according to previous studies (12, 17, 20).

## Potential predictors of vitamin D deficiency

Based on previous studies, we selected factors that may contribute to VDD (18, 21). Participants were divided into four groups based on their physiological characteristics: <18, 18–44, 45–65, and >65 years old. Race/ethnicity was classified as non-Hispanic white, non-Hispanic black Americans, Mexican Americans, other Hispanics, and other races. Body mass index (BMI) was classified into three categories: BMI <18.5, 18.5–25, and >25. The education level was divided into two groups: high-school degree and below, college degree and above. The poverty income ratio (PIR) was calculated by dividing family income by the poverty threshold in the survey year. Many studies have used it as the primary indicator of socioeconomic status (SES) (22, 23). The PIR was divided into three categories: low-income (PIR ≤ 1.3), middle-income (PIR > 1.3–3.5), and high-income (PIR > 3.5) based on previous literature (24). Based on a previous study, we also discussed the effect of sun-protective behaviors on VDD (25). We collected three sun-protective behaviors from three questionnaires—staying in the shade, wearing long-sleeved shirts, and using sunscreen. Responses for all three behaviors included “always,” “most of the time,” “sometimes,” “rarely,” and “never.” We defined sun-protective behaviors as frequent (always or most of the time), moderate (sometimes), or rare (never or rarely). Physical activity was collected through a questionnaire—how many days in the past 7 days did you perform a total of at least 60 min of exercise? We defined physical activity as rare (0 days), sometimes (1–3 days), and regular (4–7 days). Smoking and drinking behaviors were determined by a questionnaire: Have you smoked at least 100 cigarettes in your life? Have you had at least 12 alcoholic drinks in 1 year? Milk consumption was determined by a questionnaire: Have you had regular milk use five times per week with three answers, namely, “I had never been a regular milk drinker,” “I sometimes was a regular milk drinker in my life,” and “I have been a regular milk drinker for most or all of life.” Individuals with incomplete data (Label description as refused, don’t know, and missing) were pre-excluded when weighted linear regression analyses

were performed. Details on the above data can be found on the NHANES website (<http://www.cdc.gov/nchs/nhanes/>).

## Statistical analysis

All analyses and data were adjusted for the complex sampling design of NHANES using Mobile Examination Center (MEC) weights. We used percentages for categorical variables and means ± standard deviations for continuous variables. To compare the difference between the groups, we used the weighted  $\chi^2$ -test and the Wald *F*-test for categorical and continuous variables, respectively. We calculated the weighted prevalence of serum 25(OH)D levels <25, 25–50, 50–75, and >75 nmol/L by dividing the weighted number of people with different vitamin D concentrations by the weighted total number of people in the study. From 2001 to 2018, linear trend tests were used to identify trends in vitamin D status in Americans. A *P*-value of <0.05 was considered statistically significant.

Multivariate linear regression models were used to explore the predictors of serum 25(OH)D levels <25 and 50 nmol/L, controlling for a large range of variables. The following predictors were explored, including age group (<18, 18–44, 45–65, and >65 years old), sex (female and male), race/ethnicity (non-Hispanic whites, non-Hispanic blacks, Mexican Americans, other Hispanics, non-Hispanic blacks, other races), education (high-school degree and below, college degree and above), PIR (<1.3, 1.3–3.5, and >3.5), BMI (<18.5, 18.5–25, and >25 kg/m<sup>2</sup>), season (winter and summer), sun-protective behaviors, smoking behaviors, alcohol consumption, physical activity, and milk consumption. A *P*-value of <0.05 was considered statistically significant. All statistics were performed using the R software (version 4.2.0) and EmpowerStats (<http://www.empowerstats.com>).

## Results

Overall, 71,685 subjects were included in our analysis, with a mean age of 34.8 ± 24.0. In this study, 20.7% of the participants were Mexican Americans, 8.4% were other Hispanics, 38.2% were non-Hispanic whites, 23.0% were non-Hispanic blacks, and 9.7% were other races (including multiracial). 48.1% of the tests were carried out in winter, and 51.9% in summer. The characteristics of the participants are listed in [Table 1](#).

The weighted prevalence of serum 25(OH)D levels <25, 25–50, 50–75, and >75 nmol/L was 2.6% (95% CI 2.5–2.7), 22.0% (95% CI 21.7–22.3), 40.9% (95% CI 40.5–41.3), and 34.5% (95% CI 34.2–34.9) in Americans older than 1 year, respectively ([Table 2](#)).

Compared with males, females presented a higher weighted proportion of serum 25(OH)D levels <25 nmol/L (3.1% vs. 2.1%, *P* < 0.001) and 25–50 nmol/L (22.8% vs. 21.2%, *P* < 0.001),

TABLE 1 Descriptive characteristics of the participants ( $n = 71,685$ ) in the present study.

Characteristic	%	95 % CI
<b>Sex</b>		
Female	50.4	(50.3–51.1)
<b>Race/Ethnicity</b>		
Mexican American	20.7	(20.4–21.0)
Other Hispanics	8.4	(8.2–8.6)
Non-Hispanic white	38.2	(37.8–38.6)
Non-Hispanic black	23.0	(22.7–23.3)
Other race	9.7	(9.5–9.9)
<b>Age</b>		
<18	34.1	(33.8–34.5)
18–44	31.0	(30.7–31.4)
45–65	19.9	(19.8–20.0)
>65	15.0	(14.7–15.3)
<b>Education</b>		
High school degree and below	49.7	(49.2–50.2)
College degree and above	50.3	(49.8–50.9)
<b>PIR</b>		
<1.3	34.3	(33.9–34.7)
1.3–3.5	41.1	(40.8–41.5)
>3.5	24.6	(24.2–24.9)
<b>Sun-Protective Behaviors</b>		
<b>Staying in the shade</b>		
Rare	35.1	(34.5–35.8)
Sometimes	38.6	(37.9–39.3)
Frequent	26.3	(25.7–26.9)
<b>Wearing long sleeves</b>		
Rare	10.8	(10.3–11.2)
Sometimes	19.7	(19.2–20.3)
Frequent	69.5	(68.9–70.3)
<b>Using sunscreen</b>		
Rare	22.9	(22.3–23.5)
Sometimes	19.1	(18.5–19.6)
Frequent	58.0	(57.4–58.6)
<b>Season</b>		
Winter	48.1	(47.1–49.0)
Summer	51.9	(50.0–53.0)
<b>Had at least 12 alcohol drinks/1 year?</b>		
Yes	70.1	(69.6–70.5)
No	29.9	(29.5–30.4)
<b>Smoke at least 100 cigarettes in life</b>		
Yes	45.9	(45.4–46.4)
No	54.1	(53.6–54.6)
<b>Physical activity</b>		
Rare	4.5	(4.0–5.1)
Sometimes	15.1	(14.2–16.0)
Regular	80.4	(79.4–81.4)

(Continued)

TABLE 1 (Continued)

Characteristic	%	95 % CI
<b>BMI (vs. &lt;18.5)</b>		
<18.5	15.9	(15.6–16.2)
18.5–25	31.0	(30.6–31.3)
>25	53.1	(52.7–53.5)
<b>Milk consumption (vs. Rare)</b>		
Rare	22.9	(22.5–23.3)
Sometimes	35.1	(34.6–35.5)
Regular	42.0	(41.6–42.5)

PIR, poverty income ratio; BMI, body mass index.

but a lower weighted proportion of serum 25(OH)D levels 50–75 nmol/L (Table 2 and Figure 1). For ethnicity (Table 2 and Figure 2), the weighted proportion of serum 25(OH)D levels <25 and 25–50 nmol/L was lowest in non-Hispanic whites (0.9% and 13.8%,  $P < 0.001$ ) and highest in non-Hispanic blacks (11.9% and 48.5%,  $P < 0.001$ ). The weighted proportion of serum 25(OH)D levels of <25 nmol/L was lowest in aged 1–9 years (8.1% for boys and 10.1% for girls) and highest in ages between 20 and 29 years (29.1% for men and 28.1% for women). The same trend was also found in serum 25(OH)D levels of 25–50 nmol/L (Table 2 and Figure 1). Compared with summer, individuals in winter had a higher weighted proportion of serum 25(OH)D levels <25 nmol/L (4.1% vs. 1.6%,  $P < 0.001$ ) and 25–50 nmol/L (28.9% vs. 17.0%  $P < 0.001$ ), but a lower weighted proportion of serum 25(OH)D levels 50–75 nmol/L and >75 (Table 2 and Figure 1).

Table 3 and Figure 3 show the change trends of different vitamin D status in Americans from 2001 to 2018. We found a slight linear decrease in prevalence of moderate VDD [25 <25(OH)D <50 nmol/L] (coefficient =  $-0.847$ ;  $P = 0.009$ ) and VDI [50 <25(OH)D <75 nmol/L] (coefficient =  $-0.810$ ;  $P = 0.014$ ). We found a slight linear increase in vitamin D sufficient [25(OH)D >75 nmol/L] (coefficient =  $1.693$ ;  $P = 0.004$ ). However, no trend change was found in severe VDD [25(OH)D <25 nmol/L] (coefficient =  $-0.037$ ;  $P = 0.698$ ). The same trend was also found after sex-specific analysis.

## Predictors of vitamin D deficiency

When using a cut-off of 25 nmol/L, the female was an independent predictor of VDD (OR 1.39 [95% CI 1.26–1.52]). In addition, race/ethnicity was a strong independent predictor of VDD. Compared with non-Hispanic white, Mexican Americans (OR 1.81 [95% CI 1.53–2.14]), other Hispanics (OR 1.73 [95% CI 1.38–2.16]), non-Hispanic black (OR [8.59 95% CI 7.48–9.86]), other race (OR 2.93 [95% CI 2.40–3.57]) have a higher risk of VDD. Compared with individuals aged <18

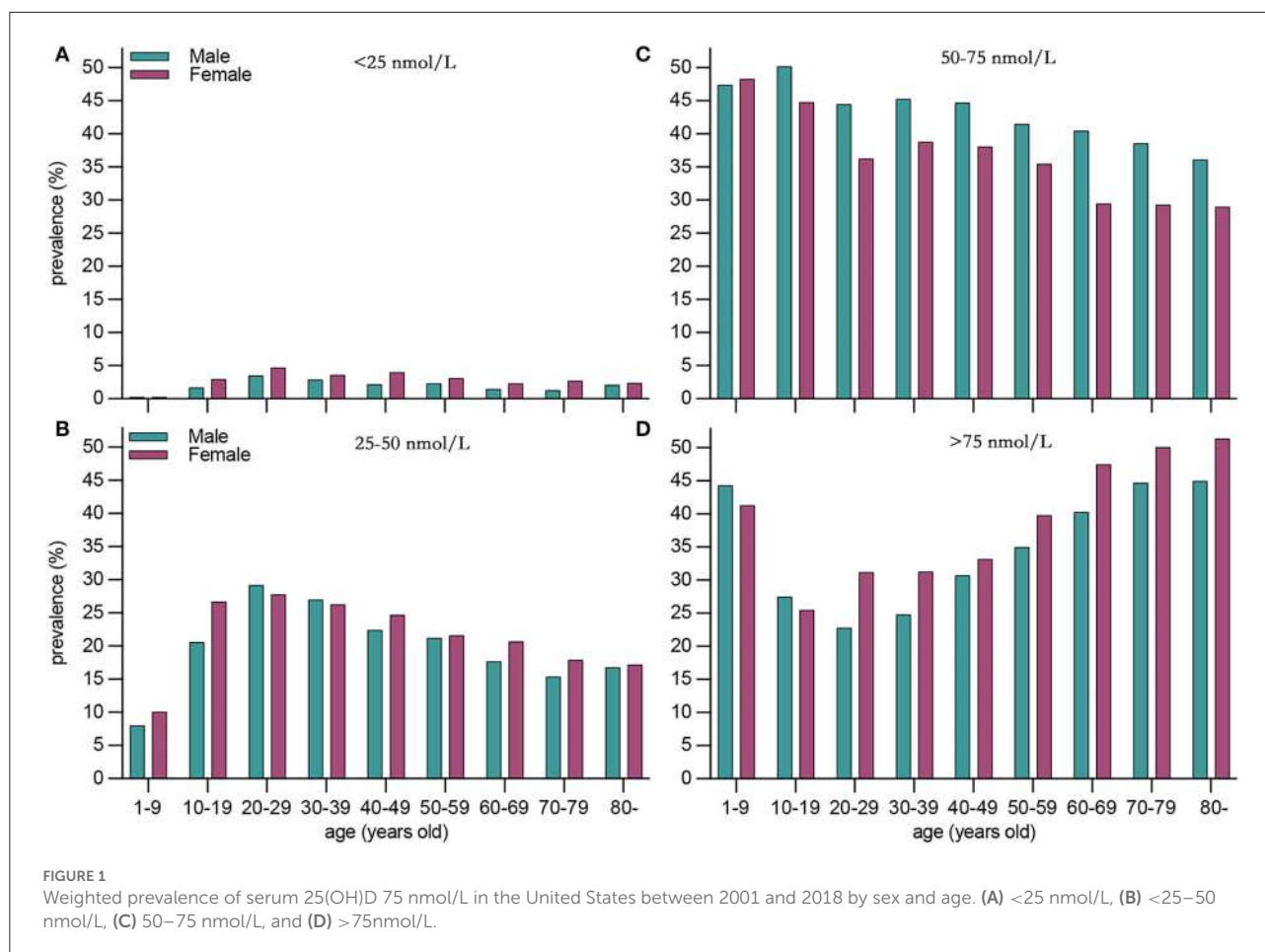
TABLE 2 Weighted prevalence of serum 25(OH)D &lt;25, 25–50, 50–75, and &gt;75 nmol/l by gender, ethnicity, age, and season.

	Sample size	<25 nmol/L	95% CI	25–50 nmol/L	95 CI	50–75 nmol/L	95% CI	>75 nmol/L	95 CI	P
All	71,685	2.6%	(2.5%, 2.7%)	22.0%	(21.7%, 22.3%)	40.9%	(40.5%, 41.3%)	34.5%	(34.2%, 34.9%)	
Gender										<0.001
Male	35,306	2.1%	(2.0%, 2.3%)	21.2%	(20.8%, 21.6%)	44.4%	(43.9%, 44.9%)	32.3%	(31.8%, 32.8%)	
Female	36,379	3.1%	(2.9%, 3.3%)	22.8%	(22.4%, 23.2%)	37.5%	(37.0%, 38.0%)	36.6%	(36.1%, 37.1%)	
Ethnicity										<0.001
Mexican American	14,822	3.2%	(2.9%, 3.5%)	35.1%	(34.3%, 35.9%)	47.5%	(46.7%, 48.3%)	14.2%	(13.7%, 14.8%)	
Other Hispanics	6,017	2.3%	(1.9%, 2.7%)	28.7%	(27.6%, 29.9%)	49.4%	(48.1%, 50.7%)	19.6%	(18.6%, 20.6%)	
Non-Hispanic white	27,411	0.9%	(0.8%, 1.0%)	13.8%	(13.4%, 14.2%)	41.1%	(40.5%, 41.7%)	44.2%	(43.6%, 44.8%)	
Non-Hispanic black	16,467	11.9%	(11.4%, 12.4%)	48.5%	(47.7%, 49.3%)	28.9%	(28.2%, 29.6%)	10.7%	(10.2%, 11.2%)	
Other race—including multi-racial	6,968	3.2%	(2.8%, 3.6%)	31.2%	(30.1%, 32.3%)	41.3%	(40.2%, 42.5%)	24.3%	(23.3%, 25.3%)	
Males—age (years)										<0.001
1–9	5,843	0.3%	(0.2%, 0.5%)	8.0%	(7.3%, 8.7%)	47.4%	(46.1%, 48.7%)	44.3%	(43.0%, 45.6%)	
10–19	8,000	1.7%	(1.4%, 2.0%)	20.0%	(19.7%, 21.5%)	50.2%	(49.1%, 51.3%)	27.5%	(26.5%, 28.5%)	
20–29	3,546	3.5%	(2.9%, 4.2%)	29.2%	(27.7%, 30.7%)	44.5%	(42.9%, 46.1%)	22.8%	(21.4%, 24.2%)	
30–39	3,500	2.9%	(2.4%, 3.5%)	27.0%	(25.6%, 28.5%)	45.3%	(43.7%, 47.0%)	24.8%	(23.4%, 26.3%)	
40–49	3,564	2.2%	(1.8%, 2.7%)	22.4%	(21.1%, 23.8%)	44.7%	(43.1%, 46.3%)	30.7%	(29.2%, 32.2%)	
50–59	3,356	2.3%	(1.8%, 2.9%)	21.2%	(19.8%, 22.6%)	41.5%	(39.9%, 43.2%)	35.0%	(33.4%, 36.6%)	
60–69	3,600	1.5%	(1.2%, 2.0%)	17.7%	(16.5%, 19.0%)	40.5%	(38.9%, 42.1%)	40.3%	(38.7%, 41.9%)	
70–79	2,427	1.3%	(0.9%, 1.9%)	15.4%	(14.0%, 16.9%)	38.6%	(36.7%, 40.6%)	44.7%	(42.7%, 46.7%)	
>80	1,470	2.1%	(1.6%, 3.1%)	16.8%	(15.0%, 18.8%)	36.1%	(33.7%, 38.6%)	45.0%	(42.5%, 47.6%)	
Females—age (years)										<0.001
1–9	5,617	0.3%	(0.2%, 0.5%)	10.1%	(9.3%, 10.9%)	48.3%	(47.0%, 49.6%)	41.3%	(40.0%, 42.6%)	
10–19	7,764	3.0%	(2.6%, 3.4%)	26.7%	(25.7%, 27.7%)	44.8%	(43.7%, 45.9%)	25.5%	(24.5%, 26.5%)	
20–29	4,106	4.7%	(4.1%, 5.4%)	27.8%	(26.4%, 29.2%)	36.3%	(34.8%, 37.8%)	31.2%	(29.8%, 32.6%)	
30–39	3,944	3.6%	(3.1%, 4.2%)	26.3%	(24.9%, 27.7%)	38.8%	(37.3%, 40.3%)	31.3%	(29.9%, 32.8%)	
40–49	3,943	4.0%	(3.4%, 4.7%)	24.7%	(23.4%, 26.1%)	38.1%	(36.6%, 39.6%)	33.2%	(31.8%, 34.7%)	
50–59	3,415	3.1%	(2.6%, 3.7%)	21.6%	(20.3%, 23.0%)	35.5%	(33.9%, 37.1%)	39.8%	(38.2%, 41.5%)	
60–69	3,656	2.3%	(1.9%, 2.8%)	20.7%	(19.4%, 22.1%)	29.5%	(28.1%, 31.0%)	47.5%	(45.9%, 49.1%)	
70–79	2,317	2.7%	(2.1%, 3.5%)	17.9%	(16.4%, 19.5%)	29.3%	(27.5%, 31.2%)	50.1%	(48.1%, 52.1%)	
>80	1,617	2.4%	(1.8%, 3.3%)	17.2%	(15.4%, 19.1%)	29.0%	(26.8%, 31.3%)	51.4%	(49.0%, 53.8%)	
Season										<0.001
Winter	34,467	4.1%	(3.9%, 4.3%)	28.9%	(28.4%, 29.4%)	40.7%	(40.2%, 41.2%)	26.3%	(25.8%, 26.8%)	
Summer	37,218	1.6%	(1.5%, 1.7%)	17.0%	(16.6%, 17.4%)	41.0%	(40.5%, 41.5%)	40.5%	(40.0%, 41.0%)	

The weighted  $\chi^2$ -test was used for categorical variables, and the Wald F-test was used for continuous variables to compare the difference between groups. P-value <0.05 was considered statistically significant. MEC weights were used. MEC, Mobile Examination Center.

years old, VDD was common in individuals aged 18–44 years old (OR 3.06 [95% CI 2.53–3.71]), 45–65 years old (OR 2.51 [95% CI 2.00–3.14]), and >65 years old (OR 2.32 [95% CI 1.80–3.00]). Being assessed during winter was also an independent predictor of VDD (OR 2.06 [95% CI 1.89–2.26]).

Moreover, lower PIR, lower education level, higher BMI, sun-protective behaviors (staying in the shade, wearing long sleeves), drinking, and lower milk consumption were also predictors of VDD. The details of predictors of VDD are listed in [Table 4](#).



## Discussion

To our knowledge, this study is the latest and most comprehensive estimate of the vitamin D status of the American population using the available data from NHANES (2001–2018). The main findings of our study were that the prevalence of serum 25(OH)D <25, 25–50, 50–75, and >75 nmol/L was 2.6%, 22.0%, 40.9%, and 34.5% in Americans aged >1 year old between 2001 and 2018. Age, sex, ethnicity, season, sun-protective behaviors, lower BMI, lower SES, drinking, and lower milk consumption were predictors of VDD.

Our results also showed that the prevalence of severe VDD [25(OH)D <25 nmol/L] had not improved significantly, and the moderate deficiency [25 <25(OH)D < 50 nmol/L], insufficiency [50 <25(OH)D < 75 nmol/L] had a mild improvement in the United States in recent years. Several potential mechanisms could explain it. First, with the increased health awareness of the general population in the United States in recent years, there has been a surge in serum 25(OH)D testing and diagnoses of VDD (26, 27). Second, healthcare providers are increasingly recommending higher doses of vitamin D

supplements than previously (28), and people have a 10-fold increase in spending on vitamin D supplements between 2001 and 2009 (29). Schleicher et al. (30) estimated the use of vitamin D supplements in Americans based on NHANES and showed a significant increase in the use of vitamin D supplements in the general population. In 2003–2004, only 0.45% of adults (>20 years) used  $\geq 1,000$  IU/d vitamin D-containing supplements. The percentage increased to 16.12% in 2013–2014. Furthermore, more and more foods in the U.S. are being fortified with vitamin D (31). Nevertheless, vitamin D status in the U.S. population has not improved significantly in recent years, which may lead to adverse consequences (32–34). More action is needed by the government and medical providers to provide effective prevention and treatment strategies for the disease.

Vitamin D deficiency is often associated with insufficient sunlight, as humans obtain most of their vitamin D through UVB radiation. A previous study reported a negative association between the serum level of 25(OH)D and distance from the equator (35). The United States is a high-latitude country where lower UVB radiation may cause a higher incidence of VDD

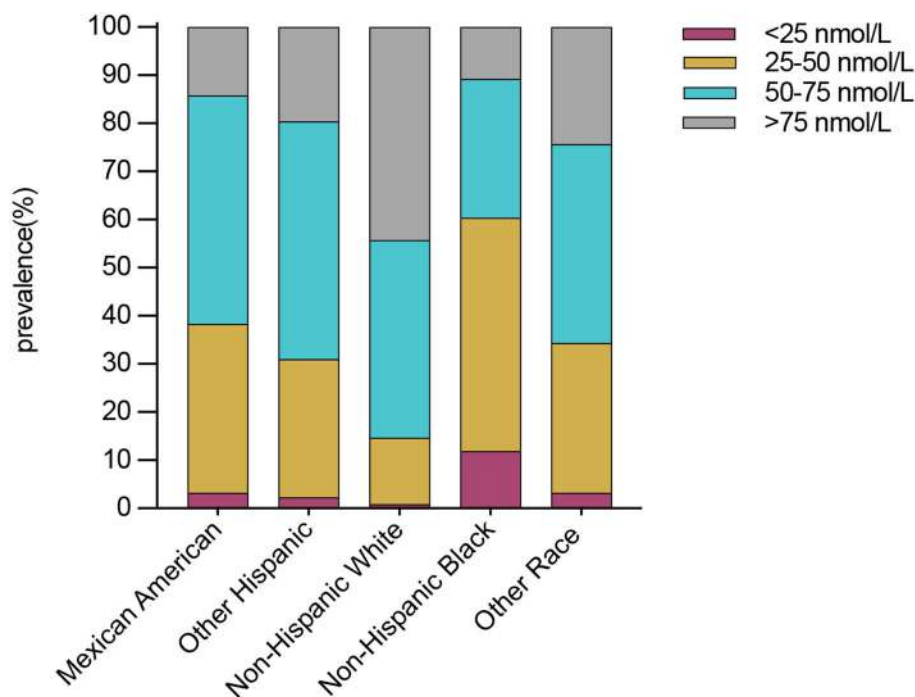


FIGURE 2

Weighted prevalence of serum 25(OH)D <25, 25–50, 50–75, and >75 nmol/L in Americans ages >1 year between 2001 and 2018 by ethnicity.

than in countries with ample sunshine (36). However, our results show that the prevalence of VDD in Americans was lower than in some low-latitude countries (13, 37), where the population has a higher direct UV-B exposure. A large meta-analysis conducted by Mogire et al. (13) reported that 18.46% and 34.22% of Africans have VDD when using a cut-off of 30 and 50 nmol/L, respectively. Pereira-Santos et al. (37) reported that 28.16% of Brazilians suffered from VDD when using a cut-off of 50 nmol/L, higher than the result of our study (24.6%). It could mean that other factors besides latitude or sun exposure may affect vitamin D status, like race/ethnicity, cultural practices, and other factors (38, 39). For instance, in the United States, food supplementation and fortification are common sources of vitamin D (40), but they are unavailable in many African and South American countries (13). Cultural practices also have an impact on vitamin D status. For example, Middle Easterners often wear veils (covering skin from sunlight), which could lead to a higher prevalence of VDD (41). Our results also show a lower prevalence of VDD [serum 25(OH)D < 50 nmol/L] in the United States than in Europe. A nationally representative study in Europe showed that 40.4% of these populations had serum 25(OH)D < 50 nmol/L (42), whereas we found a prevalence of 24.6% in the United States.

The prevalence of VDD varied greatly based on the different cut-offs, race/ethnicity, sex, age, tested season, and so on. The prevalence of serum 25(OH)D <25 and 25–50 nmol/L is

highest among non-Hispanic blacks and lowest among non-Hispanic whites, the same as the previous studies (43–45). This relationship persisted after controlling for other variables. Dark-skinned people, especially non-Hispanic blacks, have pigment melanin in their skin that can absorb sunlight, which decreases the synthesis of vitamin D (46, 47). In addition, studies have shown that lower serum 25(OH)D levels in African-Americans may be associated with obesity (48, 49). However, an article showed that non-Hispanic blacks with serum 25(OH)D below the cut-off typically lack the accompanying characteristic alterations (38). A previous community-based study showed that black Americans have similarly bioavailable 25(OH)D concentrations to white Americans, although they had lower levels of a total of 25(OH)D and a vitamin D receptor (VDR) (38). Using the same 25(OH)D cut-off to define VDD for diverse populations may not be appropriate and need further investigation.

In addition to race, the present study shows that the prevalence of serum 25(OH)D <25 and <50 nmol/L is highest in people aged 20–29 in America. After controlling for variables, our study showed that being 18–44 was also a predictor of VDD. The age-specific trends in VDD prevalence varied from study to study, but most results showed a higher prevalence in young adulthood (19, 21, 50). One explanation is that older people are more likely to use vitamin D supplements than younger people (51). Moreover, a cross-sectional study conducted in

TABLE 3 Trends of the prevalence of serum 25(OH)D &lt;25, 25–50, 50–75, and &gt;75 nmol/L in the United States from 2001 to 2018.

	Total			Males			Females		
	Sample size	Prevalence (%)	95% CI	Sample size	Prevalence (%)	95% CI	Sample size	Prevalence (%)	95% CI
<b>25(OH)D &lt;25 nmol/l</b>									
Total	71,685	2.6%	(2.5%, 2.7%)	35,306	2.1%	(2.0%, 2.3%)	36,379	3.2%	(3.0%, 3.4%)
2001–2002	7,807	2.2%	(1.9%, 2.6%)	3,782	1.3%	(1.0%, 1.7%)	4,025	3.1%	(2.6%, 3.7%)
2003–2004	8,294	3.9%	(3.5%, 4.3%)	4,095	2.5%	(2.1%, 3.0%)	4,199	5.3%	(4.7%, 6.0%)
2005–2006	8,306	1.6%	(1.4%, 1.9%)	4,048	1.3%	(1.0%, 1.7%)	4,258	2.0%	(1.6%, 2.5%)
2007–2008	6,950	3.1%	(2.7%, 3.5%)	3,481	2.4%	(2.0%, 3.0%)	3,469	3.7%	(3.1%, 4.4%)
2009–2010	8,700	3.0%	(2.7%, 3.4%)	4,321	2.3%	(1.9%, 2.8%)	4,379	3.7%	(3.2%, 4.3%)
2011–2012	7,743	2.5%	(2.2%, 2.9%)	3,879	2.4%	(2.0%, 2.9%)	3,864	2.6%	(2.1%, 3.1%)
2013–2014	8,437	2.6%	(2.3%, 3.0%)	4,134	2.4%	(2.0%, 2.9%)	4,303	2.9%	(2.4%, 3.4%)
2015–2016	8,039	2.3%	(2.0%, 2.7%)	3,956	2.2%	(1.8%, 2.7%)	4,083	2.5%	(2.1%, 3.0%)
2017–2018	7,409	2.5%	(2.2%, 2.9%)	3,610	2.2%	(1.8%, 2.7%)	3,799	2.8%	(2.3%, 3.4%)
Trend		$P = 0.698$			$P = 0.196$			$P = 2.259$	
		$\beta = -0.037$			$\beta = 0.082$			$\beta = -0.148$	
<b>25 &lt;25(OH)D &lt;50 nmol/l</b>									
Total	71,685	22.0%	(21.7%, 22.3%)	35,306	21.5%	(21.1%, 22.0%)	36,379	23.4%	(22.9%, 23.9%)
2001–2002	7,807	25.7%	(24.7%, 26.7%)	3,782	23.2%	(21.9%, 24.6%)	4,025	28.1%	(26.7%, 29.5%)
2003–2004	8,294	23.5%	(22.6%, 24.4%)	4,095	21.6%	(20.4%, 22.9%)	4,199	25.3%	(24.0%, 26.6%)
2005–2006	8,306	27.7%	(26.8%, 28.7%)	4,048	26.2%	(24.9%, 27.6%)	4,258	29.1%	(27.8%, 30.5%)
2007–2008	6,950	20.8%	(19.9%, 21.8%)	3,481	19.4%	(18.1%, 20.7%)	3,469	22.2%	(20.9%, 23.6%)
2009–2010	8,700	20.9%	(20.1%, 21.8%)	4,321	19.7%	(18.5%, 20.9%)	4,379	22.0%	(20.8%, 23.2%)
2011–2012	7,743	20.9%	(20.0%, 21.8%)	3,879	20.5%	(19.3%, 21.8%)	3,864	21.4%	(20.1%, 22.7%)
2013–2014	8,437	20.6%	(19.8%, 21.5%)	4,134	21.4%	(20.2%, 22.7%)	4,303	19.8%	(18.6%, 21.0%)
2015–2016	8,039	20.2%	(19.3%, 21.1%)	3,956	20.0%	(18.8%, 21.3%)	4,083	20.4%	(19.2%, 21.7%)
2017–2018	7,409	19.0%	(18.1%, 19.9%)	3,610	19.6%	(18.4%, 20.9%)	3,799	18.3%	(17.1%, 19.6%)
Trend		$P = 0.009$			$P = 0.108$			$P = 0.001$	
		$\beta = -0.847$			$\beta = -0.426$			$\beta = -1.222$	
<b>50 &lt;25(OH)D &lt;75 nmol/l</b>									
Total	71,685	40.9%	(40.5%, 41.3%)	35,306	44.7%	(44.2%, 45.3%)	36,379	37.8%	(37.3%, 38.3%)
2001–2002	7,807	45.1%	(44.0%, 46.2%)	3,782	48.0%	(46.4%, 49.6%)	4,025	42.3%	(40.8%, 43.8%)
2003–2004	8,294	42.4%	(41.3%, 43.5%)	4,095	45.3%	(43.8%, 46.8%)	4,199	39.7%	(38.2%, 41.2%)
2005–2006	8,306	45.7%	(44.6%, 46.8%)	4,048	47.9%	(46.4%, 49.4%)	4,258	43.6%	(42.1%, 45.1%)
2007–2008	6,950	39.9%	(38.8%, 41.1%)	3,481	44.7%	(43.1%, 46.4%)	3,469	35.3%	(33.7%, 36.9%)
2009–2010	8,700	39.6%	(38.6%, 40.6%)	4,321	44.3%	(42.8%, 45.8%)	4,379	35.1%	(33.7%, 36.5%)
2011–2012	7,743	37.6%	(36.5%, 38.7%)	3,879	40.3%	(38.8%, 41.8%)	3,864	35.0%	(33.5%, 36.5%)
2013–2014	8,437	40.7%	(39.7%, 41.8%)	4,134	44.6%	(43.1%, 46.1%)	4,303	37.0%	(35.6%, 38.5%)
2015–2016	8,039	39.1%	(38.0%, 40.2%)	3,956	43.2%	(41.7%, 44.8%)	4,083	35.1%	(33.7%, 36.6%)
2017–2018	7,409	38.5%	(37.4%, 39.6%)	3,610	42.1%	(40.5%, 43.7%)	3,799	35.2%	(33.7%, 36.7%)
Trend		$P = 0.014$			$P = 0.020$			$P = 0.021$	
		$\beta = -0.810$			$\beta = -0.682$			$\beta = -0.928$	
<b>25(OH)D &gt;75 nmol/l</b>									
Total	71,685	34.5%	(33.2%, 34.9%)	35,306	31.7%	(31.2%, 32.2%)	36,379	35.6%	(35.1%, 36.1%)
2001–2002	7,807	27.0%	(26.0%, 28.0%)	3,782	27.5%	(26.1%, 29.0%)	4,025	26.5%	(25.2%, 27.9%)

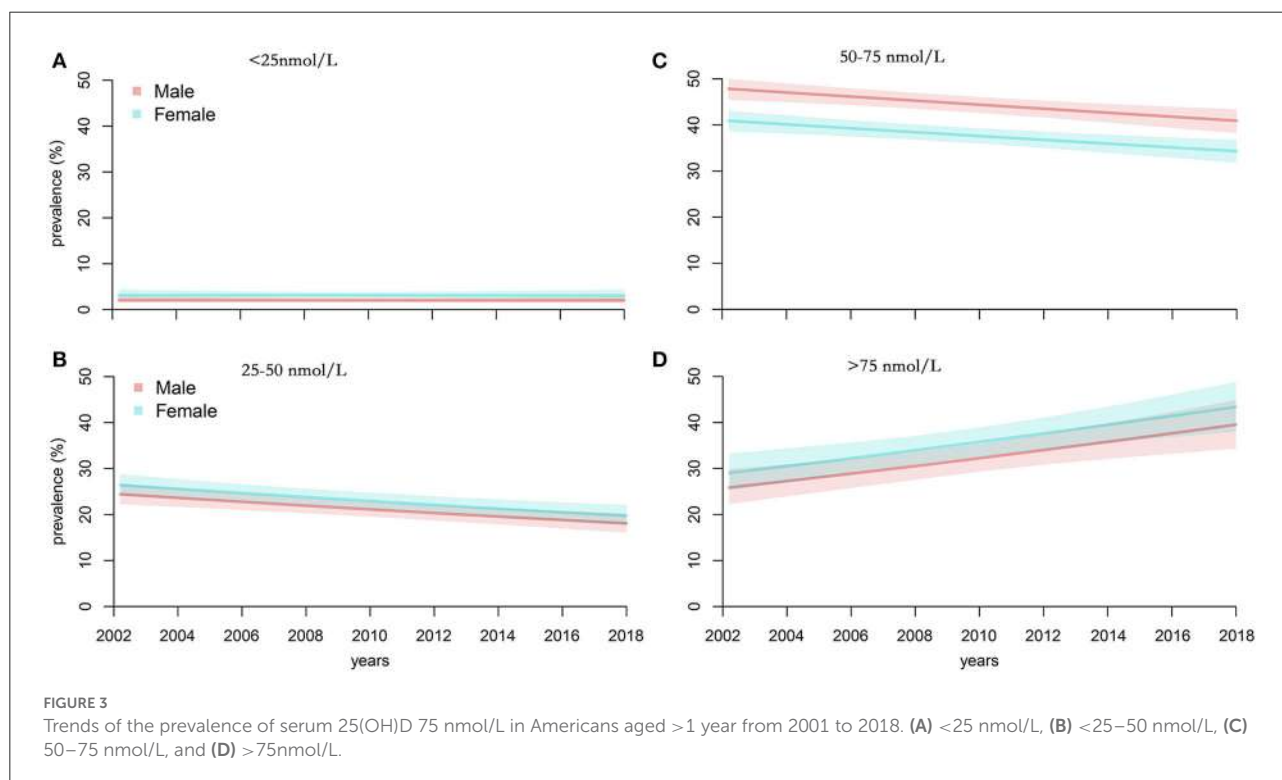
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TABLE 3 (Continued)

	Total			Males			Females		
	Sample size	Prevalence (%)	95% CI	Sample size	Prevalence (%)	95% CI	Sample size	Prevalence (%)	95% CI
2003–2004	8,294	30.2%	(29.2%, 31.2%)	4,095	30.7%	(29.3%, 32.1%)	4,199	29.7%	(28.3%, 31.1%)
2005–2006	8,306	25.0%	(24.1%, 26.0%)	4,048	24.6%	(23.3%, 26.0%)	4,258	25.3%	(24.0%, 26.6%)
2007–2008	6,950	36.2%	(35.1%, 37.3%)	3,481	33.5%	(32.0%, 35.1%)	3,469	38.8%	(37.2%, 40.4%)
2009–2010	8,700	36.4%	(35.4%, 37.4%)	4,321	33.7%	(32.3%, 35.1%)	4,379	39.1%	(37.7%, 40.6%)
2011–2012	7,743	39.0%	(37.9%, 40.1%)	3,879	36.9%	(35.4%, 38.4%)	3,864	41.1%	(39.6%, 42.7%)
2013–2014	8,437	36.1%	(35.1%, 37.1%)	4,134	31.6%	(30.2%, 33.0%)	4,303	40.4%	(38.9%, 41.9%)
2015–2016	8,039	38.4%	(37.3%, 39.5%)	3,956	34.6%	(33.1%, 36.1%)	4,083	42.1%	(40.6%, 43.6%)
2017–2018	7,409	40.0%	(38.9%, 41.1%)	3,610	36.0%	(34.5%, 37.6%)	3,799	43.7%	(42.1%, 45.3%)
Trend		$P = 0.004$			$P = 0.029$			$P = 0.001$	
		$\beta = 1.693$			$\beta = 1.052$			$\beta = 2.308$	

A linear trend test was used to identify trends in vitamin D status in Americans from 2001 to 2018. MEC weights were used. P-value <0.05 was considered statistically significant.  $\beta$ -regression coefficient, unit = % per cycle. MEC, Mobile Examination Center.



rural America shows that younger people are more likely to use sunscreen than older people (52).

In this present study, PIR and education level are considered to be the two main measures of SES (53), which have been identified as independent predictors for VDD. Individuals with low SES may have low disease awareness and may not like consuming foods high in vitamin D, such as fish and milk (54). The literature shows that low SES (low

income) may limit the potential to purchase more expensive and vitamin D-rich foods (e.g., sea fish, fish oil, fortified foods, and eggs) and do regular physical examinations to find VDD timely (55). For instance, European women with lower SES are less likely to use vitamin D supplements (56, 57). Lin et al. reported that low SES was associated with an elevated risk of VDD in women of childbearing age (58).

TABLE 4 The results of the predictors of vitamin D deficiency by weighted linear regression modeling.

	25(OH)D level <25 nmol/L			25(OH)D level <50 nmol/L		
	OR	95 % CI	P	OR	95 % CI	P
Sex (vs. men)	[Reference]			[Reference]		
Women	1.39	(1.26, 1.52)	<0.01	1.17	(1.13, 1.22)	<0.01
Race/ethnicity (vs. non-Hispanic white)	[Reference]			[Reference]		
Mexican American	1.81	(1.53, 2.14)	<0.01	2.55	(2.41, 2.69)	<0.01
Other Hispanics	1.73	(1.38, 2.16)	<0.01	2.12	(1.98, 2.28)	<0.01
Non-Hispanic black	8.59	(7.48, 9.86)	<0.01	7.16	(6.80, 7.55)	<0.01
Other race	2.93	(2.40, 3.57)	<0.01	3.06	(2.86, 3.29)	<0.01
Age (vs. <18)	[Reference]			[Reference]		
18–44	3.06	(2.53, 3.71)	<0.01	2.36	(1.86, 2.28)	<0.01
45–65	2.51	(2.00, 3.14)	<0.01	1.71	(0.88, 1.18)	<0.01
>65	2.32	(1.80, 3.00)	<0.01	1.45	(0.51, 0.75)	<0.01
Education (vs. high school degree and below)	[Reference]			[Reference]		
College degree and above	0.89	(0.80, 0.99)	0.03	0.88	(0.85, 0.93)	<0.01
PIR (vs. <1.3)	[Reference]			[Reference]		
1.3–3.5	0.89	(0.80, 0.98)	0.02	0.93	(0.89, 0.97)	<0.01
>3.5	0.68	(0.60, 0.77)	<0.01	0.74	(0.70, 0.78)	<0.01
Sun-protective behaviors						
Staying in the shade (vs. Rare)	[Reference]			[Reference]		
Sometimes	1.15	(0.94, 1.40)	0.17	1.12	(1.02, 1.22)	0.01
Frequent	1.44	(1.19, 1.75)	<0.01	1.46	(1.34, 1.60)	<0.01
Wearing long sleeves (vs. Rare)	[Reference]			[Reference]		
Sometimes	1.05	(0.88, 1.26)	0.56	1.10	(1.02, 1.21)	0.01
Frequent	1.43	(1.16, 1.77)	<0.01	1.17	(1.05, 1.30)	<0.01
Using sunscreen (vs. Rare)	[Reference]			[Reference]		
Sometimes	0.71	(0.57, 0.89)	<0.01	0.77	(0.70, 0.84)	<0.01
Frequent	0.65	(0.51, 0.82)	<0.01	0.74	(0.67, 0.81)	<0.01
Season (vs. Summer)	[Reference]			[Reference]		
Winter	2.08	(1.89, 2.26)	<0.01	0.56	(0.54, 0.59)	<0.01
Had at least 12 alcohol drinks/1 year? (vs. yes)	[Reference]			[Reference]		
No	0.83	(0.74, 0.93)	<0.01	0.91	(0.87, 0.96)	<0.01
Smoke at least 100 cigarettes in life (vs. yes)	[Reference]			[Reference]		
No	1.02	(0.91, 1.15)	0.69	1.17	(1.10, 1.24)	<0.01
Physical activity (vs. Rare)	[Reference]			[Reference]		
Sometimes	0.77	(0.35, 1.69)	0.52	0.84	(0.62, 1.14)	0.26
Regular	0.57	(0.27, 1.21)	0.14	0.58	(0.43, 0.76)	<0.01
BMI (vs. <18.5)	[Reference]			[Reference]		
18.5–25	2.21	(1.82, 2.69)	<0.01	2.21	(1.82, 2.69)	<0.01
>25	3.03	(2.49, 3.68)	<0.01	3.03	(2.49, 3.68)	<0.01
Milk consumption (vs. Rare)	[Reference]			[Reference]		
Sometimes	0.77	(0.69, 0.88)	<0.01	0.84	(0.79, 0.90)	<0.01
Regular	0.44	(0.38, 0.50)	<0.01	0.65	(0.61, 0.69)	<0.01

MEC weights were used. Adjusted for age, gender, race/ethnicity, BMI, PIR, drinking behavior, sun-protective behaviors, season, drinking behavior, smoking behavior, physical activity, and milk consumption. MEC, mobile examination center; PIR, poverty income ratio; BMI, body mass index.

Other studies have widely discussed the potential mechanisms for these connections between BMI, alcohol consumption, sun-protective behaviors, and milk consumption

with vitamin D status (59–62). Notably, the relationship between sunscreen use and vitamin D status is contrary to previous studies (63, 64). It may be due to the limitations of

the questionnaire, as regular sunscreen users tend to be more exposed to sunlight.

Our study has several obvious advantages. On the one hand, this is a large sample analysis based on the NHANES survey. The sample of this study adopted multi-layer random sampling, with high reliability and standardization of data, which can represent the general population of the United States. On the other hand, we used the currently available vitamin D data from NHANES 2001–2018 and used different cut-offs to analyze vitamin D status in Americans. Some limitations should be acknowledged. First, since this is a cross-sectional study, no causal relationship between predictors and VDD can be inferred. Second, for technical reasons, serum 25(OH)D concentrations were measured by RIA kit in 2001–2006 and LC-MS/MS method in 2007–2018, which may lead to instability of results. However, vitamin D data measured by RIA kit date were adjusted and converted to equivalent measurements by LC-MS/MS methods. Third, because vitamin D status is influenced by factors, such as season and vitamin supplement use, we were unable to assess these specific factors due to the limited information available from NHANES. Fourth, since the United States is a country with vast geographical and latitudinal differences, we could not assess them because of the limited information available from NHANES. Further studies are needed to explore it in the future.

## Conclusion

Vitamin D deficiency is still prevalent in the United States, especially in non-Hispanic blacks, women, individuals aged 20–29 years, and during the season of winter. Individuals, healthcare providers, and policymakers should take public health measures to develop and implement prevention strategies for VDD.

## Data availability statement

The datasets presented in this study can be found in online repositories. The names of the repository/repositories and accession number(s) can be found below: <https://www.cdc.gov/nchs/nhanes/>.

## References

- Holick MF. Vitamin D deficiency. *N Engl J Med.* (2007) 357:266–81. doi: 10.1056/NEJMra070553
- Bhan A, Rao AD, Rao DS. Osteomalacia as a result of vitamin D deficiency. *Rheum Dis Clin North Am.* (2012) 38:81–91, viii–ix. doi: 10.1016/j.rdc.2012.03.008
- Bouillon R, Marcocci C, Carmeliet G, Bikle D, White JH, Dawson-Hughes B, et al. Skeletal and extraskeletal actions of vitamin D: current evidence and outstanding questions. *Endocr Rev.* (2019) 40:1109–51. doi: 10.1210/er.2018-00126
- Manousaki D, Harroud A, Mitchell RE, Ross S, Forgetta V, Timpson NJ, et al. Vitamin D levels and risk of type 1 diabetes: a Mendelian randomization

## Author contributions

AC, PX, FZ, LZ, and JZ: conceptualization, project administration, and visualization. AC, PX, YM, and LZ: data curation. AC, PX, YM, ZF, FZ, LZ, and JZ: formal analysis, investigation, and writing—review and editing. AC, PX, YM, and LZ: methodology. AC, PX, and LZ: software.

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We obtained the data from the National Health and Nutrition Examination Survey (NHANES).

## Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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## Supplementary material

The Supplementary Material for this article can be found online at: <https://www.frontiersin.org/articles/10.3389/fnut.2022.965376/full#supplementary-material>

study. *PLoS Med.* (2021) 18:e1003536. doi: 10.1371/journal.pmed.1003536

5. Steinvil A, Leshem-Rubinow E, Berliner S, Justo D, Finn T, Ish-shalom M, et al. Vitamin D deficiency prevalence and cardiovascular risk in Israel. *Eur J Clin Invest.* (2011) 41:263–8. doi: 10.1111/j.1365-2362.2010.02403.x

6. Zhao J, Wang H, Zhang Z, Zhou X, Yao J, Zhang R, et al. Vitamin D deficiency as a risk factor for thyroid cancer: a meta-analysis of case-control studies. *Nutrition.* (2019) 57:5–11. doi: 10.1016/j.nut.2018.04.015

7. Autier P, Boniol M, Pizot C, Mullie P. Vitamin D status and ill health: a systematic review. *Lancet Diabetes Endocrinol.* (2014) 2:76–89. doi: 10.1016/s2213-8587(13)70165-7
8. Pugach IZ, Pugach S. Strong correlation between prevalence of severe vitamin D deficiency and population mortality rate from COVID-19 in Europe. *Wien Klin Wochenschr.* (2021) 133:403–5. doi: 10.1007/s00508-021-01833-y
9. Roth DE, Abrams SA, Aloia J, Bergeron G, Bourassa MW, Brown KH, et al. Global prevalence and disease burden of vitamin D deficiency: a roadmap for action in low- and middle-income countries. *Ann N Y Acad Sci.* (2018) 1430:44–79. doi: 10.1111/nyas.13968
10. Cashman KD. Vitamin D: dietary requirements and food fortification as a means of helping achieve adequate vitamin D status. *J Steroid Biochem Mol Biol.* (2015) 148:19–26. doi: 10.1016/j.jsbmb.2015.01.023
11. Sahota O. Understanding vitamin D deficiency. *Age Ageing.* (2014) 43:589–91. doi: 10.1093/ageing/afu104
12. Ross AC, Manson JE, Abrams SA, Aloia JF, Brannon PM, Clinton SK, et al. The 2011 report on dietary reference intakes for calcium and vitamin D from the Institute of Medicine: what clinicians need to know. *J Clin Endocrinol Metab.* (2011) 96:53–8. doi: 10.1210/jc.2010-2704
13. Mogire RM, Mutua A, Kimita W, Kamau A, Bejon P, Pettifor JM, et al. Prevalence of vitamin D deficiency in Africa: a systematic review and meta-analysis. *Lancet Glob Health.* (2020) 8:e134–42. doi: 10.1016/s2214-109x(19)30457-7
14. Ames BN, Grant WB, Willett WC. Does the high prevalence of vitamin D deficiency in African Americans contribute to health disparities? *Nutrients.* (2021) 13:499. doi: 10.3390/nu13020499
15. Yetley EA. Assessing the vitamin D status of the US population. *Am J Clin Nutr.* (2008) 88:558s–64s. doi: 10.1093/ajcn/88.2.558S
16. Sempos CT, Heijboer AC, Bikle DD, Bollerslev J, Bouillon R, Brannon PM, et al. Vitamin D assays and the definition of hypovitaminosis D: results from the First International Conference on Controversies in Vitamin D. *Br J Clin Pharmacol.* (2018) 84:2194–207. doi: 10.1111/bcp.13652
17. Holick MF, Binkley NC, Bischoff-Ferrari HA, Gordon CM, Hanley DA, Heaney RP, et al. Evaluation, treatment, and prevention of vitamin D deficiency: an Endocrine Society clinical practice guideline. *J Clin Endocrinol Metab.* (2011) 96:1911–30. doi: 10.1210/jc.2011-0385
18. Liu X, Baylin A, Levy PD. Vitamin D deficiency and insufficiency among US adults: prevalence, predictors and clinical implications. *Br J Nutr.* (2018) 119:928–36. doi: 10.1017/s0007114518000491
19. Herrick KA, Storandt RJ, Afful J, Pfeiffer CM, Schleicher RL, Gahche JJ, et al. Vitamin D status in the United States, 2011–2014. *Am J Clin Nutr.* (2019) 110:150–7. doi: 10.1093/ajcn/nqz037
20. Bouillon R. Comparative analysis of nutritional guidelines for vitamin D. *Nat Rev Endocrinol.* (2017) 13:466–79. doi: 10.1038/nrendo.2017.31
21. Malacova E, Cheang PR, Dunlop E, Sherriff JL, Lucas RM, Daly RM, et al. Prevalence and predictors of vitamin D deficiency in a nationally representative sample of adults participating in the 2011–2013 Australian Health Survey. *Br J Nutr.* (2019) 121:894–904. doi: 10.1017/s0007114519000151
22. Loucks EB, Rehkopf DH, Thurston RC, Kawachi I. Socioeconomic disparities in metabolic syndrome differ by gender: evidence from NHANES III. *Ann Epidemiol.* (2007) 17:19–26. doi: 10.1016/j.annepidem.2006.07.002
23. Gangrade N, Figueroa J, Leak TM. Socioeconomic disparities in foods/beverages and nutrients consumed by U.S. adolescents when snacking: national health and nutrition examination survey 2005–2018. *Nutrients.* (2021) 13:2530. doi: 10.3390/nu13082530
24. Ogden CL, Carroll MD, Fakhouri TH, Hales CM, Fryar CD, Li X, et al. Prevalence of obesity among youths by household income and education level of head of household - United States 2011–2014. *MMWR Morb Mortal Wkly Rep.* (2018) 67:186–9. doi: 10.15585/mmwr.mm6706a3
25. Afarideh M, Sartori-Valinotti JC, Tollefson MM. Association of sun-protective behaviors with bone mineral density and osteoporotic bone fractures in US adults. *JAMA Dermatol.* (2021) 157:1437–46. doi: 10.1001/jamadermatol.2021.4143
26. Shahangian S, Alspach TD, Astles JR, Yesupriya A, Dettwyler WK. Trends in laboratory test volumes for Medicare Part B reimbursements, 2000–2010. *Arch Pathol Lab Med.* (2014) 138:189–203. doi: 10.5858/arpa.2013-0149-OA
27. Huang KE, Milliron BJ, Davis SA, Feldman SR. Surge in US outpatient vitamin D deficiency diagnoses: National Ambulatory Medical Care Survey analysis. *South Med J.* (2014) 107:214–7. doi: 10.1097/smj.0000000000000085
28. Souberbielle JC, Body JJ, Lappe JM, Plebani M, Shoenfeld Y, Wang TJ, et al. Vitamin D and musculoskeletal health, cardiovascular disease, autoimmunity and cancer: recommendations for clinical practice. *Autoimmun Rev.* (2010) 9:709–15. doi: 10.1016/j.autrev.2010.06.009
29. Vitamin D: how 32 supplements really measure up. *Consum Rep.* (2013) 78:48.
30. Schleicher RL, Sternberg MR, Potischman N, Gahche JJ, Storandt RJ, Maw KL, et al. Supplemental vitamin D increased serum total 25-hydroxyvitamin D in the US adult population during 2007–2014. *J Nutr.* (2021) 151:2446–54. doi: 10.1093/jn/nxab147
31. Calvo MS, Whiting SJ. Survey of current vitamin D food fortification practices in the United States and Canada. *J Steroid Biochem Mol Biol.* (2013) 136:211–3. doi: 10.1016/j.jsbmb.2012.09.034
32. Hashemipour S, Larijani B, Adibi H, Javadi E, Sedaghat M, Pajouhi M, et al. Vitamin D deficiency and causative factors in the population of Tehran. *BMC Public Health.* (2004) 4:38. doi: 10.1186/1471-2458-4-38
33. Mallah EM, Hamad MF, Elmanaseer MA, Qinna NA, Idkaidek NM, Arafat TA, et al. Plasma concentrations of 25-hydroxyvitamin D among Jordanians: effect of biological and habitual factors on vitamin D status. *BMC Clin Pathol.* (2011) 11:8. doi: 10.1186/1472-6890-11-8
34. Song HR, Kweon SS, Choi JS, Rhee JA, Lee YH, Nam HS, et al. High prevalence of vitamin D deficiency in adults aged 50 years and older in Gwangju, Korea: the Dong-gu Study. *J Korean Med Sci.* (2014) 29:149–52. doi: 10.3346/jkms.2014.29.1.149
35. Durazo-Arvizu RA, Camacho P, Bovet P, Forrester T, Lambert EV, Plange-Rhule J, et al. 25-Hydroxyvitamin D in African-origin populations at varying latitudes challenges the construct of a physiologic norm. *Am J Clin Nutr.* (2014) 100:908–14. doi: 10.3945/ajcn.113.066605
36. Engelsen O. The relationship between ultraviolet radiation exposure and vitamin D status. *Nutrients.* (2010) 2:482–95. doi: 10.3390/nu2050482
37. Pereira-Santos M, Santos J, Carvalho GQ, Santos DBD, Oliveira AM. Epidemiology of vitamin D insufficiency and deficiency in a population in a sunny country: geospatial meta-analysis in Brazil. *Crit Rev Food Sci Nutr.* (2019) 59:2102–9. doi: 10.1080/10408398.2018.1437711
38. Powe CE, Evans MK, Wenger J, Zonderman AB, Berg AH, Nalls M, et al. Vitamin D-binding protein and vitamin D status of black Americans and white Americans. *N Engl J Med.* (2013) 369:1991–2000. doi: 10.1056/NEJMoa1306357
39. Marzban M, Kalantarhormozi M, Mahmudpour M, Ostovar A, Keshmiri S, Darabi AH, et al. Prevalence of vitamin D deficiency and its associated risk factors among rural population of the northern part of the Persian Gulf. *BMC Endocr Disord.* (2021) 21:219. doi: 10.1186/s12902-021-00877-5
40. Calvo MS, Whiting SJ, Barton CN. Vitamin D fortification in the United States and Canada: current status and data needs. *Am J Clin Nutr.* (2004) 80:1710s–6s. doi: 10.1093/ajcn/80.6.1710S
41. Ahmed WA, Al-Murrani W, Kuri V, Rees GA. Vitamin D intake and other risk factors for vitamin D insufficiency in Middle Eastern people living in the UK: a comparison of cultural and ethnic groups. *Ecol Food Nutr.* (2013) 52:191–202. doi: 10.1080/03670244.2012.706105
42. Cashman KD, Dowling KG, Škrabáková Z, Gonzalez-Gross M, Valtueña J, De Henauw S, et al. Vitamin D deficiency in Europe: pandemic? *Am J Clin Nutr.* (2016) 103:1033–44. doi: 10.3945/ajcn.115.12.0873
43. Whiting SJ, Langlois KA, Vatanparast H, Greene-Finestone LS. The vitamin D status of Canadians relative to the 2011 Dietary Reference Intakes: an examination in children and adults with and without supplement use. *Am J Clin Nutr.* (2011) 94:128–35. doi: 10.3945/ajcn.111.013268
44. Kift R, Berry JL, Vail A, Durkin MT, Rhodes LE, Webb AR. Lifestyle factors including less cutaneous sun exposure contribute to starkly lower vitamin D levels in U.K. South Asians compared with the white population. *Br J Dermatol.* (2013) 169:1272–8. doi: 10.1111/bjd.12518
45. Looker AC, Johnson CL, Lacher DA, Pfeiffer CM, Schleicher RL, Sempos CT. Vitamin D status: United States, 2001–2006. *NCHS Data Brief.* (2011) 1–8.
46. Looker AC. Body fat and vitamin D status in black versus white women. *J Clin Endocrinol Metab.* (2005) 90:635–40. doi: 10.1210/jc.2004-1765
47. Matsuoka LY, Wortsman J, Haddad JG, Kolm P, Hollis BW. Racial pigmentation and the cutaneous synthesis of vitamin D. *Arch Dermatol.* (1991) 127:536–8.
48. Parikh SJ, Edelman M, Uwaifo GI, Freedman RJ, Semega-Janneh M, Reynolds J, et al. The relationship between obesity and serum 1,25-dihydroxy vitamin D concentrations in healthy adults. *J Clin Endocrinol Metab.* (2004) 89:1196–9. doi: 10.1210/jc.2003-031398

49. Rajakumar K, de las Heras J, Chen TC, Lee S, Holick MF, Arslanian SA. Vitamin D status, adiposity, and lipids in black American and Caucasian children. *J Clin Endocrinol Metab.* (2011) 96:1560–7. doi: 10.1210/jc.2010-2388
50. Duarte C, Carvalheiro H, Rodrigues AM, Dias SS, Marques A, Santiago T, et al. Prevalence of vitamin D deficiency and its predictors in the Portuguese population: a nationwide population-based study. *Arch Osteoporos.* (2020) 15:36. doi: 10.1007/s11657-020-0695-x
51. Kantor ED, Rehm CD, Du M, White E, Giovannucci EL. Trends in dietary supplement use among US adults from 1999–2012. *JAMA.* (2016) 316:1464–74. doi: 10.1001/jama.2016.14403
52. Luong J, Davis RE, Chandra A, White AE, Harber ID, Engel AA, et al. A cross-sectional survey of prevalence and correlates of sunscreen use among a rural Tri-State Appalachian population. *Arch Dermatol Res.* (2021) 313:295–8. doi: 10.1007/s00403-020-02072-3
53. Du Y, Zhao LJ, Xu Q, Wu KH, Deng HW. Socioeconomic status and bone mineral density in adults by race/ethnicity and gender: the Louisiana osteoporosis study. *Osteoporos Int.* (2017) 28:1699–709. doi: 10.1007/s00198-017-3951-1
54. Forrest KY, Stuhldreher WL. Prevalence and correlates of vitamin D deficiency in US adults. *Nutr Res.* (2011) 31:48–54. doi: 10.1016/j.nutres.2010.12.001
55. Islam MZ, Lamberg-Allardt C, Kärkkäinen M, Outila T, Salamatullah Q, Shamim AA. Vitamin D deficiency: a concern in premenopausal Bangladeshi women of two socio-economic groups in rural and urban region. *Eur J Clin Nutr.* (2002) 56:51–6. doi: 10.1038/sj.ejcn.1601284
56. Novaković R, Cavelaers A, Geelen A, Nikolić M, Altaba II, Viñas BR, et al. Socio-economic determinants of micronutrient intake and status in Europe: a systematic review. *Public Health Nutr.* (2014) 17:1031–45. doi: 10.1017/s1368980013001341
57. McCartney DM, Younger KM, Walsh J, O'Neill M, Sheridan C, Kearney JM. Socio-economic differences in food group and nutrient intakes among young women in Ireland. *Br J Nutr.* (2013) 110:2084–97. doi: 10.1017/s0007114513001463
58. Lin S, Jiang L, Zhang Y, Chai J, Li J, Song X, et al. Socioeconomic status and vitamin D deficiency among women of childbearing age: a population-based, case-control study in rural northern China. *BMJ Open.* (2021) 11:e042227. doi: 10.1136/bmjopen-2020-042227
59. Daly RM, Gagnon C, Lu ZX, Magliano DJ, Dunstan DW, Sikaris KA, et al. Prevalence of vitamin D deficiency and its determinants in Australian adults aged 25 years and older: a national, population-based study. *Clin Endocrinol (Oxf).* (2012) 77:26–35. doi: 10.1111/j.1365-2265.2011.04320.x
60. Skaaby T, Husemoen LL, Thuesen BH, Pisinger C, Hannemann A, Jørgensen T, et al. Longitudinal associations between lifestyle and vitamin D: a general population study with repeated vitamin D measurements. *Endocrine.* (2016) 51:342–50. doi: 10.1007/s12020-015-0641-7
61. Yuan L, Ni J. The association between tobacco smoke exposure and vitamin D levels among US general population, 2001–2014: temporal variation and inequalities in population susceptibility. *Environ Sci Pollut Res Int.* (2022) 29:32773–87. doi: 10.1007/s11356-021-17905-5
62. Zhang J, Cao ZB. Exercise: a possibly effective way to improve vitamin D nutritional status. *Nutrients.* (2022) 14:2652. doi: 10.3390/nu14132652
63. de Gruijl FR, Webb AR, Rhodes LE. Everyday sunscreen use may compromise vitamin D in temperate climates. *Br J Dermatol.* (2020) 182:1312–3. doi: 10.1111/bjd.18811
64. Norval M, Wulf HC. Does chronic sunscreen use reduce vitamin D production to insufficient levels? *Br J Dermatol.* (2009) 161:732–6. doi: 10.1111/j.1365-2133.2009.09332.x